### Is restenosis a benign event?

Tullio Palmerini
University of Bologna
Italy

### **Conflict of interest**

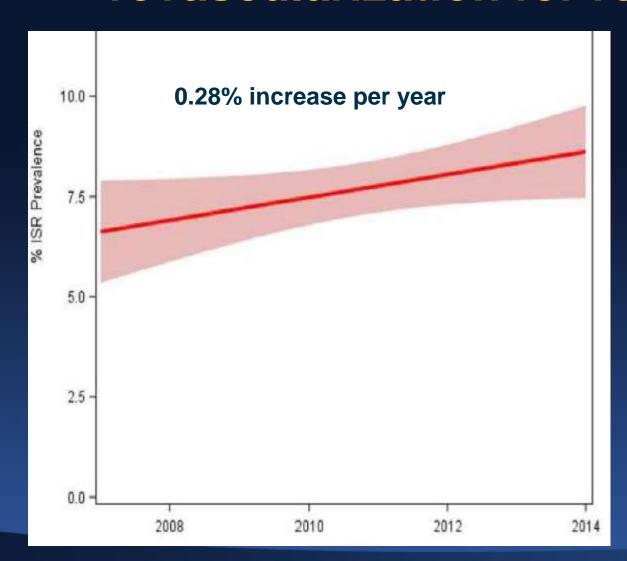
None

# In-stent restenosis: conception or misconception?

- 1) ISR usually presents with SIHD
- 2) ISR is not associated with MACE at fup
- 3) Rates of repeat revascularization for ISR declining
- 4) We need ISR in clinical trials to increase the nuber of expected events and lower the sample size



# Proportion of patients undergoing revascularization for restenosis

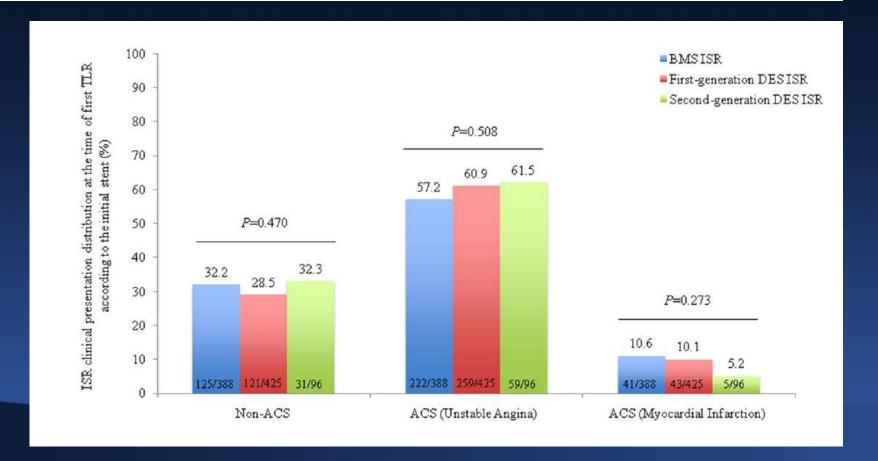


National VA Program 65,443 patients 6,872 revsc for restenosis

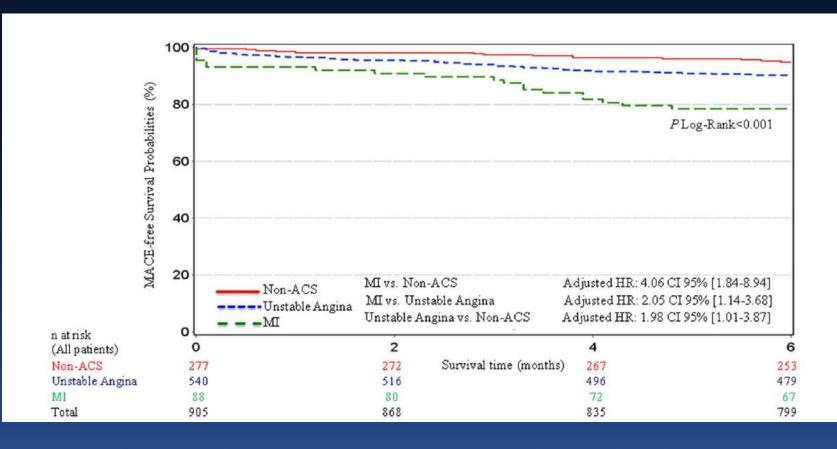
#### Clinical Presentation and Outcomes of Coronary In-Stent Restenosis Across 3-Stent Generations

Marco A. Magalhaes, MD; Sa'ar Minha, MD; Fang Chen, PhD; Rebecca Torguson, MPH;
 Al Fazir Omar, MD; Joshua P. Loh, MBBS; Ricardo O. Escarcega, MD;
 Michael J. Lipinski, MD, PhD; Nevin C. Baker, DO; Hironori Kitabata, MD, PhD;
 Hideaki Ota, MD; William O. Suddath, MD; Lowell F. Satler, MD; Augusto D. Pichard, MD;
 Ron Waksman, MD

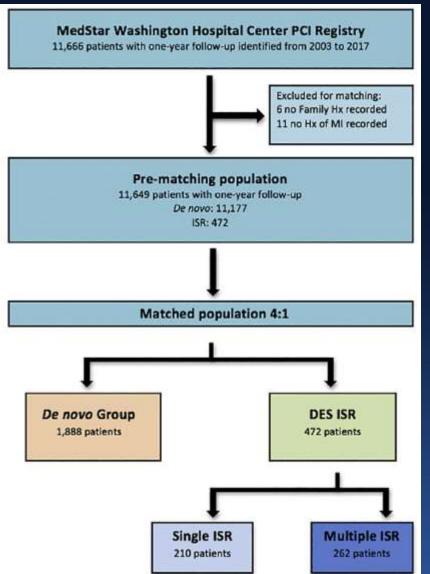
Circ Cv Int 2014

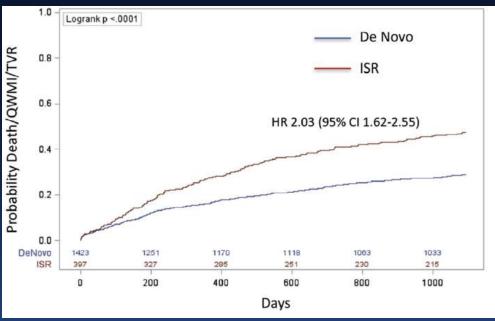


# MACE according to ISR clinical presentation



### ISR versus de novo lesions



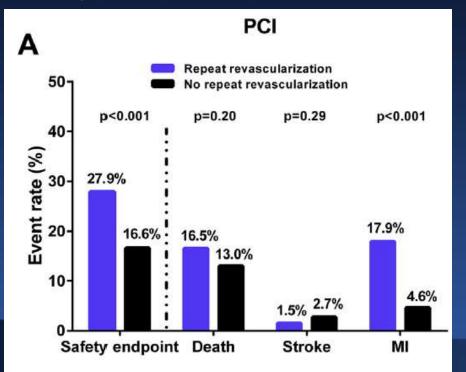


Incidence, Characteristics, Predictors, and Outcomes of Repeat Revascularization After Percutaneous Coronary Intervention and Coronary Artery Bypass Grafting

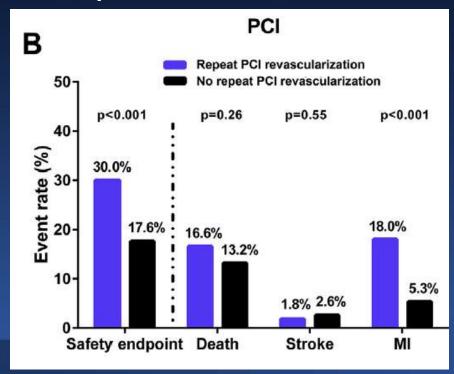
The SYNTAX Trial at 5 Years

Parasca el; JACC Int 2016

#### Any repeat revascularization



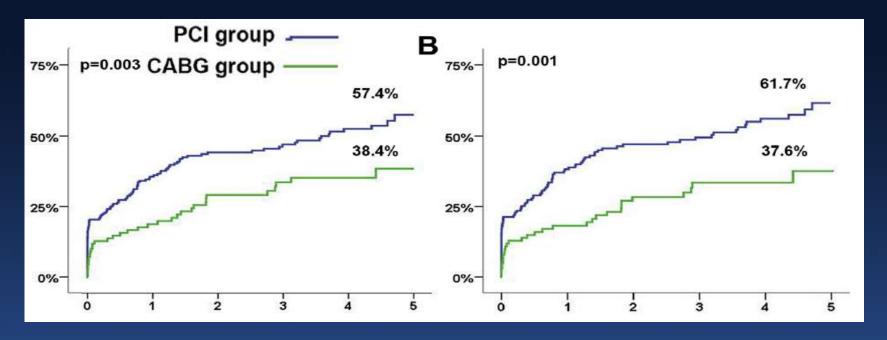
#### Repeat PCI revascularization



### Death, stroke or MI

Any repeat revascularization

Repeat PCI revascularization



#### What is known

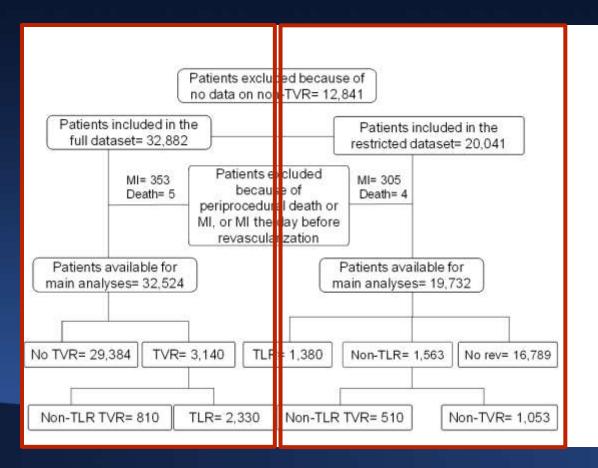
- 1)ISR may present with ACS or MI
- 2) Clinical presentation of ISR has a relevant clinical impact
- 3) ISR has a worse outcome than de novo lesions
- 4) TLR is associated with increased rates of death MI, or stroke compared to patients with no repeat revasc

#### What is not known

- 1) Prognostic implication of ISR not associated with ACS
- 2) Prognostic impplication of uncomplicated repeat revasc
- 3) Prognostic implication of different type of repeat revasc
- 4) Association with mortality?

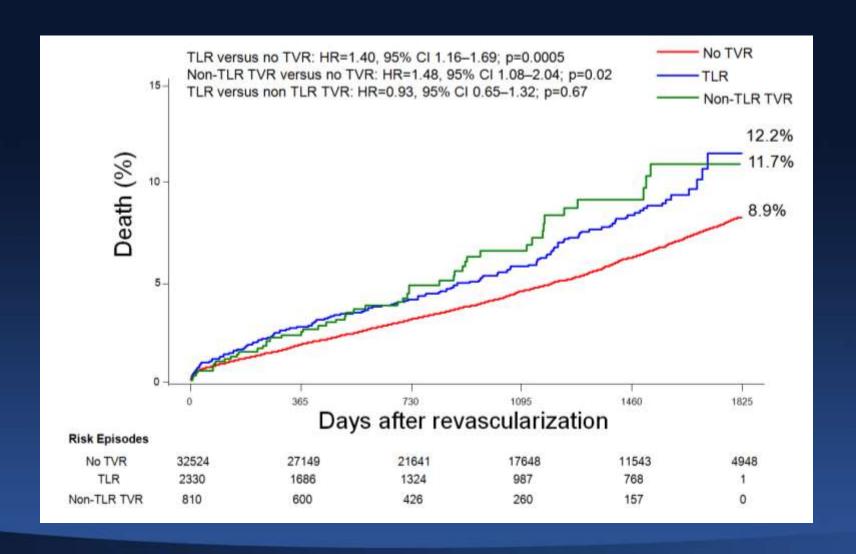
# Mortality Following Non-emergent, Uncomplicated Target Lesion Revascularization After PCI: An Individual Patient Data Pooled Analysis of 21 Randomized Trials and 32,524 Patients

Tullio Palmerini, Diego Della Riva, Giuseppe Biondi-Zoccai, Martin B. Leon, Patrick W. Serruys, Pieter C. Smits, Clemens von Birgelen, Ori Ben-Yehuda, Philippe Généreux, Antonio G. Bruno, Paul Jenkins, Gregg W. Stone



JACC Int 2018; in press

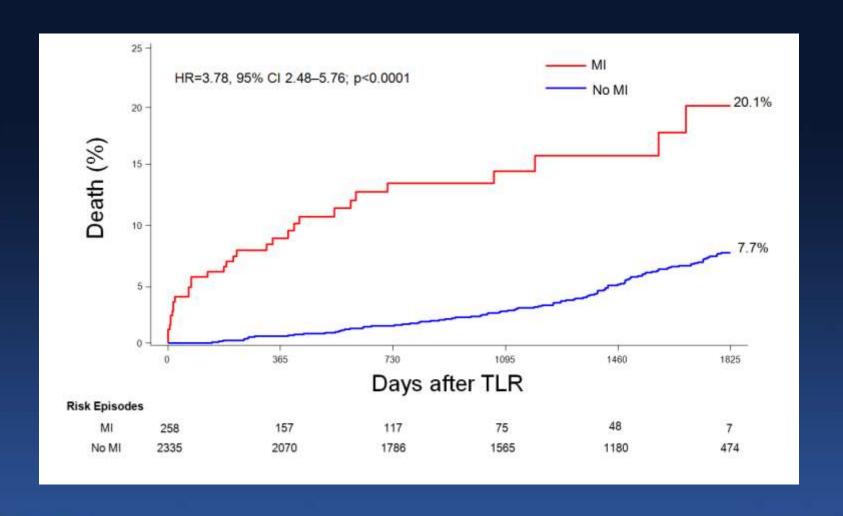
### Simon Makuch analysis of mortality



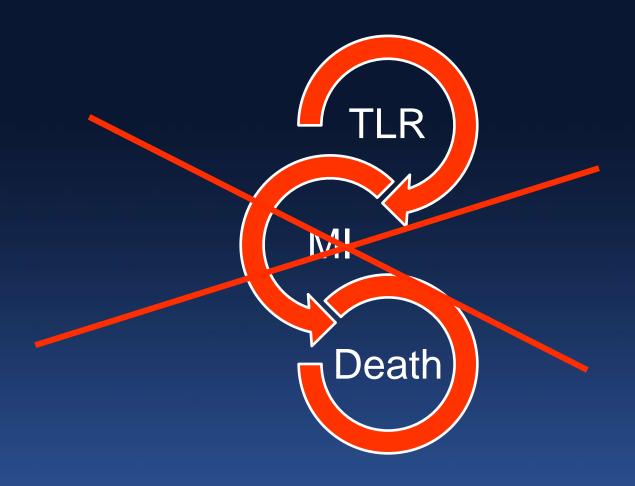
## Independent predictors of mortality

	HR (95% CI)	P value	
TLR	1.23 (1.04-1.45)	0.02	
Non-TLR TVR	1.23 (0.83-1.82)	0.31	
MI or ST during fup	4.26 (3.16-5.74)	<0.0001	
Age (per 1 year)	1.07 (1.07-1.08)	<0.0001	
Diabetes	1.60 (1.46-1.76)	<0.0001	
Male sex	1.16 (1.08-1.25)	<0.0001	
Previous CABG	1.35 (1.21-1.52)	<0.0001	
Previous MI	1.32 (1.23-1.40)	<0.0001	
Presentation with MI	1.47 (1.23-1.75)	<0.0001	
Palmerini et al; JACC Int 2018 in press			

## Mortality after TLR in patients with vs without MI

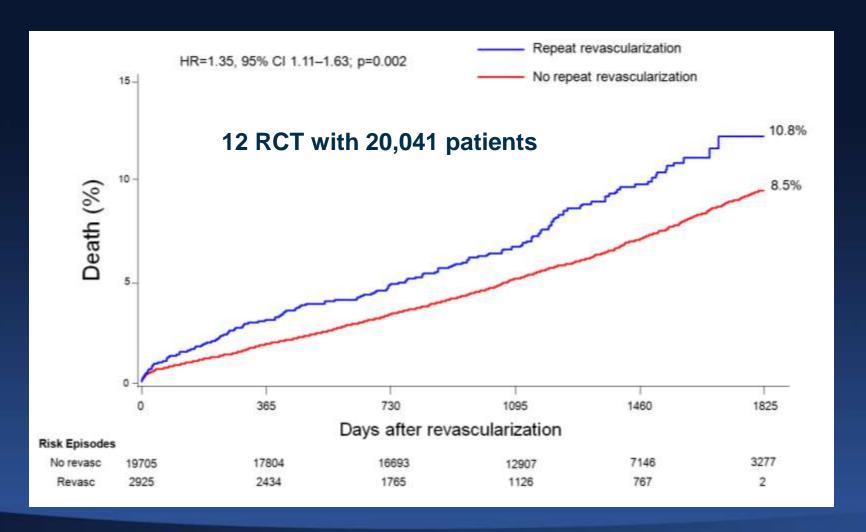


### TLR-MI-Death: a mechanistic link?

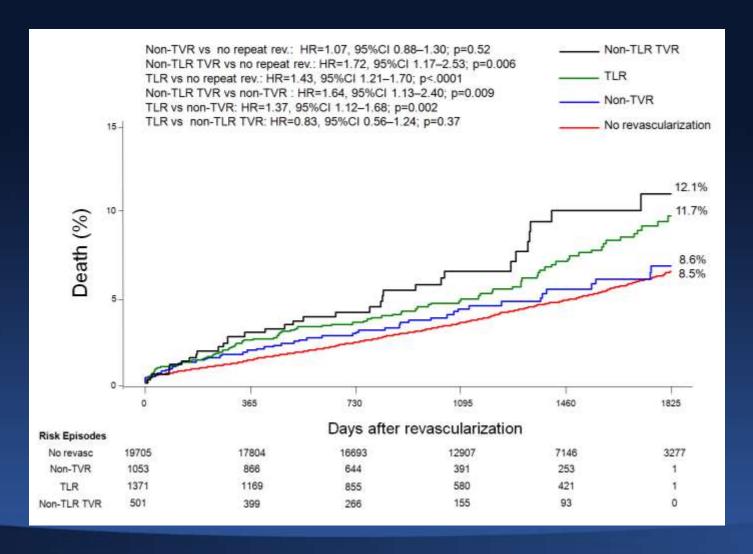


HR=1.25, 95% CI 1.04-1.50, p=0.02

# Mortality in patients with versus without any repeat revascularization



## Mortality according to the type of repeat revascularization



## Independent predictors of mortality

	HR (95% CI)	P value
TLR	1.33 (1.08-1.64)	0.02
Non-TLR	1.18 (0.90-1.55)	0.18
MI or ST	3.26 (2.27-4.68)	<0.0001
Age (per 1 year)	1.08 (1.07-1.08)	<0.0001
Diabetes	1.50 (1.39-1.61)	<0.0001
Male sex	1.20 (1.12-1.29)	<0.0001
Previous CABG	1.36 (1.19-1.56)	<0.0001
Previous MI	1.33 (1.23-1.44)	<0.0001
Presentation with MI	1.40 (1.12-1.75)	0.003
Palmerini et al; JACC Int 2018 in press		

#### **Conclusions**

- Although DES have significantly reduce the risk of restenosis compared with BMS, repeat revascularization procedure for restenosis in the last 10 years have not declined.
- In contrast to common perception, restenosis and TLR are not benign entities as they may be associated with increased rates of mortality.
- Reducing restenosis and TLR rates may therefore translate in better survival after percutaneous coronary revascularization.